PATENT COOPERATION TREATY

PCT

INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference		of Transmittal of International Search Report
UGVU-EPP/GM1	ACTION (Form PCT/ISA	√220) as well as, where applicable, item 5 below.
International application No.	International filing date (day/month/year)	(Earliest) Priority Date (day/month/year)
PCT/IB 03/00280	10/01/2003	
Applicant		•
CENTRO DE INVESTIGACIONES	Y ESTUDIOS AVANZADOS	
This later street County Bornet has been		
according to Article 18. A copy is being tra	n prepared by this International Searching Adansmitted to the International Bureau.	uthonty and is transmitted to the applicant
This International Search Report consists	of a total of6 sheets.	
X It is also accompanied by	a copy of each prior art document cited in th	ais report.
Basis of the report		
a. With regard to the language, the	international search was carried out on the bess otherwise indicated under this item.	pasis of the international application in the
the international search w Authority (Rule 23.1(b)).	as carried out on the basis of a translation o	f the international application furnished to this
b. With regard to any nucleotide an was carried out on the basis of the		international application, the international search
i —	e sequence listing : anal application in written form.	
filed together with the inte	rnational application in computer readable for	orm.
furnished subsequently to	this Authority in written form.	
	this Authority in computer readble form.	
	osequently furnished written sequence listing s filed has been furnished.	does not go beyond the disclosure in the
. —		n is identical to the written sequence listing has been
2. Certain claims were fou	nd unsearchable (See Box I).	
3. X Unity of invention is lac	king (see Box II).	
4 1454		
4. With regard to the title , X the text is approved as su	hmitted by the applicant	·
ı =	thed by this Authority to read as follows:	
	,	
. 0		
5 450		
5. With regard to the abstract, The text is approved as su	shmitted by the applicant	
the text has been establis		ority as it appears in Box III. The applicant may, report, submit comments to this Authority.
6. The figure of the drawings to be publ	ished with the abstract is Figure No.	13
X as suggested by the appli	cant.	None of the figures.
because the applicant fail	ed to suggest a figure.	
because this figure better	characterizes the invention.	

INTERNATIONAL SEARCH REPORT

International Application No PCT/IB 03/00280

A. CLASSIFICATION OF SUBJECT MATTER
IPC 7 C07K14/14 A61K39/15 A61K38/16

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

 $\begin{array}{ll} \mbox{Minimum documentation searched (classification system followed by classification symbols)} \\ \mbox{IPC 7} & \mbox{C07K} & \mbox{A61K} \end{array}$

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

BIOSIS, SEQUENCE SEARCH, EPO-Internal, MEDLINE, WPI Data, PAJ, CHEM ABS Data

J. DOCUMENTS CONSIDERED TO BE RELEVANT

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Α	WO 89 06971 A (UNIV LELAND STANFORD JUNIOR) 10 August 1989 (1989-08-10)	1-7, 18-21, 23-25, 27,29-42
	the whole document	
A	DATABASE EBI Database accession no. P12473 XP002260536 100% identical to Seq-ID-N° 1, 3 in a 776aa overlap; 100% identical to Seq-ID-N° 2 in a 231 aa overlap abstract	1-7, 18-21, 23-25, 27,29-42
	-/	
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Further documents are listed in the continuation of box C.	Patent family members are listed in annex.
Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier document but published on or after the international filing date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed	 *T* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention *X* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone *Y* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. *&* document member of the same patent family
Date of the actual completion of the international search Date of mailing of the international search report	
26 November 2003	02/12/2003
Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL – 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nt, Fax: (+31-70) 340-3016	Authorized officer Seranski, P

INTERNATIONAL SEARCH REPORT

International Application No PCT/IB 03/00280

vol. 75, no. 16, August 2001 (2001-08), pages 7339-7350, XP002260531	C.(Continu	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	<u> </u>
monomeric recombinant rotavirus VP4 yields 18-21, 23-25, 27,29-42 no ligomeric VP5* core" 23-25, 23-25, 23-25, 27,29-42 no ligomeric VP5* core" 23-25, 27,29-42 no ligomeric VP5* core" 23-25, 27,29-42 no ligomeric VP5* core" 23-25, 27,29-42 no ligomeric VP5* core 23-25, 27,29-42 abstract 1-7, 18-21, 23-25, 27,29-42 abstract 1-7,	Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No
A B DATABASE EBI Database accession no. Q91HI9 abstract A TAFAZOLI FARIDEH ET AL: "NSP4 enterotoxin of rotavirus induces paracellular leakage in polarized epithelial cells" JOUNNAL OF VIROLOGY, vol. 75, no. 3, February 2001 (2001-02), pages 1540-1546, XP002260532 ISSN: 0022-538X page 1541, column 2 -page 1542, column 1 abstract A DICKMAN KATHLEEN G ET AL: "Rotavirus alters paracellular permeability and energy metabolism in Caco-2 cells" AMERICAN JOURNAL OF PHYSIOLOGY, vol. 279, no. 4 Part 1, October 2000 (2000-10), pages G757-G766, XP002260533 ISSN: 0002-9513 the whole document A TIHOVA MARIANA ET AL: "Localization of membrane permeabilization and receptor binding sites on the VP4 hemagglutinin of rotavirus: Implications for cell entry" JOURNAL OF MOLECULAR BIOLOGY, vol. 314, no. 5, 14 December 2001 (2001-12-14), pages 985-992, XP002260534 ISSN: 0022-2836 the whole document T ZARATE SELENE ET AL: "Interaction of rotaviruses with Hsc70 during cell entry is mediated by VP5." JOURNAL OF VIROLOGY, vol. 77, no. 13, July 2003 (2003-07), pages 7254-7260, XP00920526	A	monomeric recombinant rotavirus VP4 yields an oligomeric VP5* core" JOURNAL OF VIROLOGY, vol. 75, no. 16, August 2001 (2001-08), pages 7339-7350, XP002260531 ISSN: 0022-538X	18-21,
A TAFAZOLI FARIDEH ET AL: "NSP4 enterotoxin of rotavirus induces paracellular leakage in polarized epithelial cells" 23-25, JOURNAL OF VIROLOGY, vol. 75, no. 3, February 2001 (2001-02), pages 1540-1546, XP002260532 ISSN: 0022-538X page 1541, column 2 -page 1542, column 1 abstract	A	& DATABASE EBI Database accession no. Q91HI9	18-21,
Of rotavirus induces paracellular leakage 18-21, 1n polarized epithelial cells" 23-25, 23-25, 23-25, 27,29-37 23-25, 27,29-37 23-25, 27,29-37 23-25, 27,29-37 23-25, 27,29-37 23-25, 27,29-37 23-25, 27,29-37 23-25, 27,29-37 23-25, 27,29-37 23-25, 27,29-37 23-25, 27,29-37 23-25, 27,29-37 23-25,		abstract	
alters paracellular permeability and energy metabolism in Caco-2 cells" AMERICAN JOURNAL OF PHYSIOLOGY, vol. 279, no. 4 Part 1, October 2000 (2000-10), pages G757-G766, XP002260533 ISSN: 0002-9513 the whole document A TIHOVA MARIANA ET AL: "Localization of membrane permeabilization and receptor binding sites on the VP4 hemagglutinin of 23-25, rotavirus: Implications for cell entry" JOURNAL OF MOLECULAR BIOLOGY, vol. 314, no. 5, 14 December 2001 (2001-12-14), pages 985-992, XP002260534 ISSN: 0022-2836 the whole document T ZARATE SELENE ET AL: "Interaction of rotaviruses with Hsc70 during cell entry is mediated by VP5." JOURNAL OF VIROLOGY, vol. 77, no. 13, July 2003 (2003-07), pages 7254-7260, XP009020526	Α	of rotavirus induces paracellular leakage in polarized epithelial cells" JOURNAL OF VIROLOGY, vol. 75, no. 3, February 2001 (2001-02), pages 1540-1546, XP002260532 ISSN: 0022-538X page 1541, column 2 -page 1542, column 1	18-21,
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		rotaviruses with Hsc70 during cell entry is mediated by VP5." JOURNAL OF VIROLOGY, vol. 77, no. 13, July 2003 (2003-07), pages 7254-7260, XP009020526	18-21,

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FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. Claims: 1-7, 18-21,23-25, 27, 29-37

Use of Rotavirus protein VP4 and its variants, use of antibodies against VP4, pharmaceutical compositions comprising VP4 or its variants; Methods of Treatment comprising the use of VP4 or its variants and methods for determining regions pf VP4, VP8 that enhance or modulate the opening of the paracellular pathway.

2. Claim: 38

An isolated polypeptide with Seq-ID-N°3

3. Claim: 39

An isolated polypeptide with Seq-ID-N°4

4. Claim: 40

An isolated polypeptide with Seq-ID-N°5

5. Claim: 41

An isolated polypeptide with Seq-ID-N°6

6. Claim: 42

An isolated polypeptide with Seq-ID-N°7

International application No. PCT/IB 03/00280

INTERNATIONAL SEARCH REPORT

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1 Chams Nos tecause they relate to subject matter not required to be searched by this Authority, namely:
2. Claims Nos.: tecause they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. Chirms Nos tecause they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
see additional sheet
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. X As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

INTERNATIONAL SEARCH REPORT

Information on patent family members		PC1	PCT/IB 03/00280		
Patent document cited in search report	Publication date		Patent family member(s)		Publication date
WO 8906971	A 10-08-1989	WO	8906971 A	l	10-08-1989
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